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| FORM | | | Filing Date | August 1, 2001 |
| (to be used for all correspondence after initial filing) | | First Named Inventor | Nicolau | |
| | | Art Unit | 1626 | |
| | | Examiner Name | Anderson, R.L. | |
| Total Number of Pages in This Submission 7 | | 7 | Attorney Docket Number | GMX-005.01 |
| ENCLOSURES (check all that apply) | | | | |
| Fee Transmittal Form | | ☐ Drawin | g(s) | After Allowance Communication to Group |
| Fee Attached | | Licensi | ing-related Papers | Appeal Communication to Board of Appeals and Interferences |
| Amendment / Reply | | Petition | 1 | Appeal Communication to Group (Appeal Notice, Brief, Reply Brief) |
| After Final | | Petition to Convert to a Provisional Application | | Proprietary Information |
| Affidavits/declaration(s) | | Power of Attorney, Revocation Change of Correspondence Address | | Status Letter |
| Extension of Time Request | | Terminal Disclaimer | | Other Enclosure(s) (please identify below): |
| Express Abandonment Request | | _ | st for Refund umber of CD(s) | Response to Non-Responsive Reply (6 pages); Return postcard |
| ☐ Information Disclosure Statement | | | | |
| Certified Copy of Priority Document(s) | | Rema | rks | |
| Response to Missing Parts/ Incomplete Application | | | | |
| Response to Missing Parts under 37 CFR 1.52 or 1.53 | | | | |
| | SIGNA | TURE OF | APPLICANT, ATTORNEY, O | R AGENT |
| Firm or Individual name | Michael DiVerdi | | | |
| Signature | Michael J. D. Verdi | | | |
| Date June 17, 2004 | | | | |
| | C | ERTIFICA | TE OF TRANSMISSION/MAI | LING |
| | correspondence i | is being facs | simile transmitted to the USPTO an an envelope addressed to: Co | LING or deposited with the United States Post ommissioner for Patents, P.O. Box 145 |

Signature

Date

June 17, 2004

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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

1626 Whu

In re Application of: Nicolau, Y.C. et al.

Serial No: 09/920,310

Filed: August 1, 2001

For: Ammonium Salts Of Hemoglobin Allosteric Effectors, and Uses Thereof

Attorney Docket No.: GMX-005.01

Examiner: Anderson, R. L.

Group Art Unit: 1626

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Certificate of Mailing

I hereby certify that this "Response to Non-Responsive Reply" is being deposited with the U. S. Postal Service as First Class Mail with sufficient postage on the date set forth below in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

6-14.09

Date of Signature and Mail Deposit

Steve Pereira

Response to Non-Responsive Reply

Dear Examiner Anderson:

In a Non-Responsive Reply dated May 19, 2004, it was stated that the Applicants' selected species, made in a reply dated February 25, 2004, to a restriction requirement dated January 23, 2004, does not fall within elected Group I. The Applicants respectfully traverse this contention.

It was stated that the elected group requires an aliphatic ammonium cation and that the Applicants' election of a species having cyclohexyl ammonium cations does not satisfy this requirement because aliphatic by definition means an open chain hydrocarbon and not a cyclic one. The Applicants respectfully traverse this definition of aliphatic and respectfully submit that aliphatic includes both cyclic and open chain hydrocarbons.

To support their position, the Applicants have included pages 34 and 78 from Streitwieser and Heathcock's "Introduction to Organic Chemistry," third edition, 1985, which defines aliphatic as excluding aromatic compounds, but including both cyclic and open chain hydrocarbons.

Therefore, based on what one of ordinary skill in the art would consider aliphatic, as supported by text book definitions, the Applicants respectfully submit that their elected species comprising cyclohexyl ammonium cations falls within the elected group, and that the election made on February 25, 2004, was fully responsive.

<u>Fees</u>

The Applicants believe no fee is due in connection with the filing of this paper.

Nevertheless, the Director is hereby authorized to charge any required fee to our Deposit Account, 06-1448.

Conclusion

The Applicants believe that they have responded fully to the Office communication dated May 19, 2004. However, if a telephone conversation with Applicant's Agent would expedite prosecution of the above-identified application, the Examiner is urged to contact the undersigned.

By:

Respectfully submitted, Foley Hoag LLP

Michael J. DiVerdi, PhD

Reg. No. 51,620

Agent for Applicants

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Telephone: (617) 832-1000 Telecopier: (617) 832-7000

Date: <u>June 17, 2004</u>

INTRODUCTION to ORGANIC CHEMISTRY

Andrew Streitwicser, Jr. Clayton H. Heathcock

THIRD EDITION

Chap. 3 Organic

Structures

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AND COPY SO Some Ethers

$$CH_3$$
-O- CH_3 CH_3 -O- CH_3 H_2 C- CH_2 H_2 C CH_3

The carbon-oxygen double bond, the carbonyl group, is found in aldehydes and ketones. When the carbonyl group is bonded to an OH group, it becomes a carboxy group. Compounds containing this functional group are called carboxylic acids.

Some Aldehydes

Some Carboxylic Acids

Table 3.1 lists a number of the important functional groups. The structures and names of these groups should be committed to memory. They form an essential part of the language of organic chemistry. In our subsequent studies we will develop the chemistry of the individual functional groups in terms of structural and electronic theory, nomenclature (names), physical properties, the preparation from other functional groups, and the characteristic reactions that produce other groups.

Interconversions of functional groups constitute a large proportion of organic chemistry. After the individual groups have been studied, the effect of one group on another can be considered, for the organic chemistry of compounds with more than one functional group is not simply the sum of the parts. Groups affect each other, sometimes in complex ways. One of the reasons for studying the theory of organic chemistry is that the mutual interactions of functional groups can be understood.

The aromatic ring in Table 3.1 is written with three carbon-carbon double bonds. Nevertheless, we shall see later (Chapters 20 and 22) that compounds containing this ring system differ substantially in their chemistry from the alkenes. Compounds containing this ring system are known collectively as aromatic compounds. Compounds with no aromatic ring are known as aliphatic compounds.

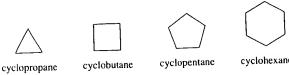
EXERCISE 3.2 Using R = ethyl, write structural and condensed formulas for one example each of an alkene, an alkyne, an alcohol, an ether, an aldehyde, a ketone, a carboxylic acid, an amine, and a nitrile.

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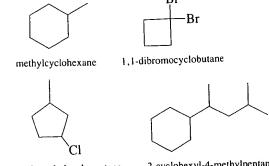
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Chap. 5 Alkanes

Cycloalkanes are usually symbolized by simple geometric figures in which a carbon atom with its appropriate number of attached hydrogens is understood to be present at each apex. Thus, the four smallest cycloalkanes are depicted as



Simple substituted cycloalkanes are depicted by the appropriate geometric figure, with attached substituent groups.



1-chloro-3-methylcyclopentane

2-cyclohexyl-4-methylpentane

The alkanes and cycloalkanes are the parent structures in the general class of aliphatic compounds. Most of the chemistry of cycloalkanes is similar to that of the alkanes. There are some differences in stability and in their conformations, which will be discussed in Sections 5.6 and 7.7, respectively.

EXERCISE 5.5 Using simple geometric figures and line structures, depict the following compounds. Compare your structures with complete structural representations.

(a) 1,1,3-trimethylcyclohexane

(b) 3-cyclopentylpentane

(c) 1-chloro-4-chloromethylcyclohexane

(d) 1,1,2,2-tetramethylcyclopropane

5.5 Heats of Formation

The heat of formation of a compound from its elements in their standard states is a thermodynamic property with considerable use in organic chemistry. This quantity, symbolized ΔH_f° , is defined as the enthalpy of the reaction of elements in their standard states to form the compound. The standard state of each element is generally the most stable state of that element at 25°C and 1 atm pressure. The standard state of carbon is taken as the graphite form, whereas those of hydrogen and oxygen are H2 and O2 gases, respectively. By definition, ΔH_f° for an element in its standard state is zero. The standard heat of formation of butane is -30.36 ± 0.16 kcal mole⁻¹ and that of 2methylpropane is -32.41 ± 0.13 kcal mole⁻¹.

4 C (graphite) + 5 H₂(g) =
$$n$$
-C₄H₁₀(g)
$$\Delta H^{\circ} \equiv \Delta H_{f}^{\circ}(n$$
-C₄H₁₀) = -30.36 kcal mole⁻¹
4 C (graphite) + 5 H₂(g) = (CH₃)₃CH(g)
$$\Delta H^{\circ} \equiv \Delta H_{f}^{\circ}(i$$
-C₄H₁₀) = -32.41 kcal mole⁻¹